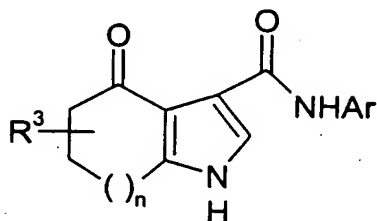
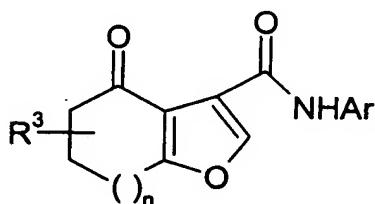


We claim:

1. A method of preparing a compound of the formula:



comprising reacting a compound of the formula:



with an excess of ammonia source in a reaction inert solvent at an elevated temperature until reaction is complete;

wherein Ar is phenyl or heterocycle, said phenyl or heterocycle being substituted with -O-(CH₂)_m-NR¹R², -O(CH₂)_lC(O)OR⁴, -CH(NR⁷R⁸)CH₃, -CH₂CH(NR⁵R⁶)CH₃, or OH, and said phenyl or heterocycle being optionally substituted with one or two groups selected from C₁ - C₆ alkoxy, C₁ - C₆ alkyl, C₂ - C₆ alkenyl, C₁ - C₆ perfluoroalkyl, F, Cl, and Br, wherein:

R¹, R³, R⁴, R⁵ and R⁷ are independently selected from hydrogen and C₁ - C₆ alkyl;

R², R⁶, and R⁸ are independently selected from nitrogen protecting groups;

m and l are integers independently selected from 1 to 6; and

n is an integer from 0 to 2.

2. The method of claim 1 wherein Ar is phenyl substituted with said one or two groups.

3. The method of claim 1 wherein said nitrogen protecting group is -C(O)C₁-C₈ alkoxy.

4. The method of claim 1 wherein said nitrogen protecting group is benzyloxycarbonyl, fluorenyloxycarbonyl, acetyl, trifluoroacetyl, chloroacetyl, benzoyl, t-butyloxycarbonyl, or benzyl.

5. The method of claim 1 wherein said compound of formula I is selected from the group consisting of

Methyl-(1-{4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl}-ethyl)-carbamate acid tert-butyl ester;

[2-(2-Fluoro-4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy)-ethyl]-propyl-carbamic acid tert-butyl ester;

Butyl-(2-{5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-pyridin-2-yloxy}-ethyl)-carbamic acid tert-butyl ester;

5 4-Oxo-4,5,6,7,8-hexahydro-cyclohepta[b]pyrrole-3-carboxylic acid (2-fluoro-4-hydroxy-phenyl)-amide;

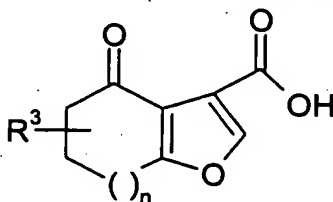
(1-Methyl-2-{4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl}-ethyl)-carbamic acid tert-butyl ester;

10 (2-{4-[(4-Oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-ethyl)-propyl-carbamic acid tert-butyl ester; and

{2-Fluoro-5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-acetic acid ethyl ester.

6. A method according to claim 1 further wherein said compound of formula II is prepared by

15 (a) reacting a compound of the formula



with an excess of an acid chloride or anhydride in a reaction inert solvent containing an excess of an acid acceptor until reaction is complete; and

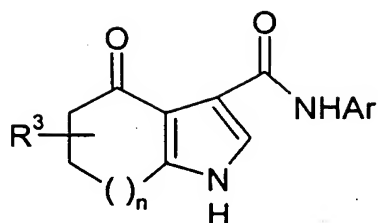
20 (b) adding an equivalent amount of $\text{NH}_2\text{-Ar}$ to the solution of step (a) and holding until reaction is complete.

7. The method of claim 6 wherein said acid chloride is ethylchloroformate.

8. The method according to claim 1 which further comprises removing said nitrogen protecting group.

25 9. The method according to claim 5 which further comprises removing said nitrogen protecting group.

10. A compound of the following formula:



wherein Ar is phenyl or heterocycle, said phenyl or heterocycle being substituted with -O-(CH₂)_m-NR¹R², -O(CH₂)C(O)OR⁴, -CH(NR⁷R⁸)CH₃, -CH₂CH(NR⁵R⁶)CH₃, or OH, and said phenyl or heterocycle being optionally substituted with one or two groups selected from C₁ - C₆ alkoxy, C₁ - C₆ alkyl, C₂ - C₆ alkenyl, C₁ - C₆ perfluoroalkyl, F, Cl, and Br, wherein:

R¹, R³, R⁴, R⁵ and R⁷ are independently selected from hydrogen and C₁ - C₆ alkyl;

R², R⁶, and R⁸ are independently selected from nitrogen protecting groups;

m and l are integers independently selected from 1 to 6; and

n is an integer from 0 to 2.

11. A compound of claim 10 selected from the group consisting of:

Methyl-(1-{4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl}-ethyl)-carbamic acid tert-butyl ester;

{2-(2-Fluoro-4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy)-ethyl}-propyl-carbamic acid tert-butyl ester;

Butyl-(2-{5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-pyridin-2-yloxy}-ethyl)-carbamic acid tert-butyl ester;

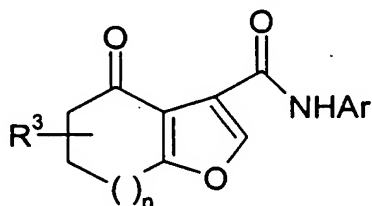
4-Oxo-4,5,6,7,8-hexahydro-cyclohepta[b]pyrrole-3-carboxylic acid (2-fluoro-4-hydroxy-phenyl)-amide;

(1-Methyl-2-{4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl}-ethyl)-carbamic acid tert-butyl ester;

(2-{4-[(4-Oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-ethyl)-propyl-carbamic acid tert-butyl ester; and

{2-Fluoro-5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-acetic acid ethyl ester.

12. A compound of the following formula:



wherein Ar is phenyl or heterocycle, said phenyl or heterocycle being substituted with $-O-(CH_2)_m-NR^1R^2$, $-O(CH_2)_lC(O)OR^4$, $-CH(NR^7R^8)CH_3$, $-CH_2CH(NR^5R^6)CH_3$, or OH, and said phenyl or heterocycle being optionally substituted with one or two groups selected from $C_1 - C_6$ alkoxy, $C_1 - C_6$ alkyl, $C_2 - C_6$ alkenyl, $C_1 - C_6$ perfluoroalkyl, F, Cl, and Br, wherein:

- 5 R^1 , R^3 , R^4 , R^5 and R^7 are independently selected from hydrogen and $C_1 - C_6$ alkyl;
 R^2 , R^6 , and R^8 are independently selected from nitrogen protecting groups;
 m and l are integers independently selected from 1 to 6; and
 n is an integer from 0 to 2.

13. The compound of claim 12 selected from the group consisting of:
- 10 Methyl-(1-{4-[(4-oxo-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-amino]-phenyl}-ethyl)-carbamic acid tert-butyl ester;
 [2-(2-Fluoro-4-[(4-oxo-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-amino]-phenoxy)-ethyl]-propyl-carbamic acid tert-butyl ester;
 Butyl-(2-{5-[(4-oxo-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-amino]-pyridin-2-yloxy}-ethyl)-carbamic acid tert-butyl ester;
- 15 4-Oxo-4,5,6,7,8-hexahydro-cyclohepta[b]furan-3-carboxylic acid (2-fluoro-4-hydroxy-phenyl)-amide;
 (1-Methyl-2-{4-[(4-oxo-4,5,6,7-tetrahydrobenzofuran-3-carbonyl)-amino]-phenyl}-ethyl)-carbamic acid tert-butyl ester;
- 20 (2-{4-[(4-Oxo-4,5,6,7-tetrahydrobenzofuran-3-carbonyl)-amino]-phenoxy}-ethyl)-propyl-carbamic acid tert-butyl ester; and
 {2-Fluoro-5-[(4-oxo-4,5,6,7-tetrahydrobenzofuran-3-carbonyl)-amino]-phenoxy}-acetic acid ethyl ester.

10008294: 120301